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Sheet 1 of 8

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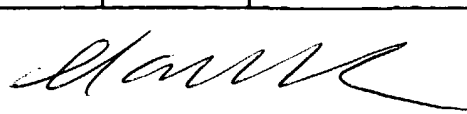
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Mahanthappa Nagesh K.Filing Date
October 14, 1999Group Art Unit
1646**U.S. PATENT DOCUMENTS**

EXAMINER INITIAL	DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
W	BJ 5,759,811	06/02/98	Epstein et al.	435	69.1	11/13/96
	BK 5,223,408	06/29/93	Goeddel et al.	435	69.3	07/11/91
	BL 4,456,687	06/26/84	Howard Green	435	241	12/01/80
	BM 5,789,543	08/04/98	Ingham et al.	530	350	12/30/93
	BN 5,844,079	12/01/98	Ingham et al.	530	350	12/14/94
	BO 5,585,087	12/17/96	Lustig et al.	424	9.2	06/08/94
	BP 5,837,538	11/17/98	Scott et al.	435	325	10/06/95
	BQ 5,747,507	05/05/98	Ikegaki et al.	514	312	08/10/93
	BR 5,643,915	07/01/97	Andrulis, Jr. et al.	514	279	06/06/95
	BS 5,519,035	05/21/96	Maiese et al.	514	309	07/02/93

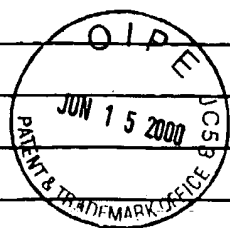
FOREIGN PATENT DOCUMENTS

	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	Translation	
						YES	NO
W	BT WO 90/02809	3/22/90	PCT	C 12P	21/00		
	BU WO 92/15679	9/17/92	PCT	C 12N	15/10		
	BV WO 94/28016	12/08/94	PCT	C 07K	13/00		
	BW WO 95/23223	08/31/95	PCT	C 12N	15/00		
	BX WO 95/18856	07/13/95	PCT	C 12N	15/12		
	BY WO 96/09806	04/04/96	PCT				
	BZ WO 96/11260	04/18/96	PCT	C 12N	5/00		
	CA WO 96/16668	06/06/96	PCT	A 61K	38/17		
	CB WO 96/17924	06/13/96	PCT	C 12N			
	CC WO 97/11095	03/27/97	PCT	C 07K	14/475		
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	CI WO 98/30576	07/16/98	PCT	C 07K	1/100		
	CJ WO 98/35020	08/13/98	PCT	C 12N	5/00		

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✓	CK	WO 99/00117	01/07/99	PCT	A 61K	31/00		
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	CN	WO 99/10004	03/04/99	PCT	A 61K	38/00		
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	CX	Angier, N., "Biologists find key genes that shape patterning of embryos", <i>New York Times</i> , Jan. 11, 1994, C-1.
	CY	Basler, K. and G. Struhl, "Compartment boundaries and the control of <i>Drosophila</i> limb pattern by Hedgehog protein", <i>Nature</i> 368 :208-214 (1994).
	CZ	Basler, K. et al., "Control of cell pattern in the neural tube: Regulation of cell differentiation by <i>dorsalin-1</i> , a novel TGF β family member", <i>Cell</i> 73 :687-702 (1993).
	DA	Bass, S. et al., "Hormone phage: An enrichment method for variant proteins with altered binding properties", <i>PROTEINS: Structure, Function, and Genetics</i> 8 :309-314 (1990).
	DB	Bejsovec, A. and E. Wieschaus, "Segment polarity gene interactions modulate epidermal patterning in <i>Drosophila</i> embryos", <i>Development</i> 119 :501-517 (1993).
	DC	Bienz, M., "Homeotic genes and positional signalling in the <i>Drosophila</i> viscera", <i>TIG</i> 10 :22-26 (Jan. 1994).
	DD	Bitgood, M. and A. McMahon, "Hedgehog and Bmp genes are coexpressed at many diverse sites of cell-cell interaction in the mouse embryo", <i>Dev. Biol.</i> 172 (1):126-138 (1995).
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	DG	Brockes, J., "We may not have a morphogen", <i>Nature</i> 350 :15 (1991).
	DH	Bumcrot, D. A. et al., "Proteolytic processing yields two secreted forms of sonic hedgehog", <i>Mol. Cell. Biol.</i> 15 (4):2294-2303 (April 1995).
	DI	Bumcrot, D. A. and A. McMahon, "Sonic hedgehog: Making the gradient", <i>Chem. Biol.</i> 3 (1):13-16 (Jan 1996).
	DJ	Bumcrot, D. A. and A. McMahon, "Somite differentiation. Sonic signals somites", <i>Curr. Biol.</i> 5 (6):612-614 (June 1995).
	DK	Charité, J. et al., "Ectopic expression of <i>Hoxb-8</i> causes duplication of the ZPA in the forelimb and homeotic transformation of axial structures", <i>Cell</i> 78 :589-601 (1994).
	DL	Coffman, et al., "Xotch, the <i>Xenopus</i> homolog of <i>Drosophila</i> notch", <i>Science</i> 249 :1438-1441 (1990).

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DN	Curry, et al., "Sequence analysis reveals homology between two proteins of the flagellar radial spoke", <i>Mol. Cell. Biol.</i> 12 :3967-3977 (1992).
DO	Davidson, E. H., "How embryos work: a comparative view of diverse modes of cell fate specification", <i>Develop.</i> 108 :365-389 (1990).
DP	Davis, A. P. and M. R. Capecchi, "Axial homeosis and appendicular skeleton defects in mice with a targeted disruption of <i>hoxd-1</i> ", <i>Devel.</i> 120 :2187-2198 (1994).
DQ	Dickinson, W., "Molecules and morphology: Where's the homology", <i>TIG</i> 11 (4):119-120 (1995).
DR	Dingemans, M. A. et al., "The expression of liver-specific genes within rat embryonic hepatocytes is a discontinuous process", <i>Differentiation</i> 56 :153-162 (1994).
DS	Dollé, P. et al., "Coordinate expression of the murine <i>Hox-5</i> complex homeobox-containing genes during limb pattern formation", <i>Nature</i> 342 :767-772 (1989).
DT	Dollé, P. et al., "Disruption of the <i>Hoxd-13</i> gene induces localized heterochrony leading to mice with neotenic limbs", <i>Cell</i> 75 :431-441 (1993).
DU	Echelard, Y. et al., "Sonic hedgehog, a member of a family of putative signaling molecules, is implicated in the regulation of CNS polarity", <i>Cell</i> 75 :1417-1430 (1993).
DV	Ekker, S. et al., "Distinct expression and shared activities of members of the hedgehog gene family of <i>xenopus laevis</i> ", <i>Devel.</i> 121 (8):2337-2347 (Aug. 1995).
DW	Ericson, J. et al., "Sonic hedgehog induces the differentiation of ventral forebrain neurons: a common signal for ventral patterning within the neural tube", <i>Cell</i> 81 (5):747-756 (June 1995).
DX	Ettelaie, C. et al., "The effect of lipid peroxidation and lipolysis on the ability of lipoproteins to influence thromboplastin activity", <i>Biochim. Biophys. Acta.</i> 1257 (1):25-30 (June 1995).
DY	Fahmer, K. et al., "Transcription of <i>H-2</i> and <i>Qa</i> genes in embryonic and adult mice", <i>EMBO J.</i> 6 :1265-1271 (1987).
DZ	Fallon, J. F. et al., "FGF-2: Apical ectodermal ridge growth signal for chick limb development", <i>Science</i> 264 :104-107 (1994).
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EC	Forbes, A. J. et al., "Genetic analysis of <i>hedgehog</i> signaling in the <i>Drosophila</i> embryo", <i>Devel.</i> 119 (Suppl.):115-124 (1993).
ED	Francis, P. H. et al., "Bone morphogenetic proteins and a signaling pathway that controls patterning in the developing chick limb", <i>Devel.</i> 120 :209-218 (1994).
EE	Gallop, M. et al., "Applications of combinatorial technologies to drug discovery. 1. Background and peptide combinatorial libraries", <i>J. Med. Chem.</i> 37 (9):1233-1251 (1994).
EF	Gérard, M. et al., "Structure and activity of regulatory elements involved in the activation of the <i>Hoxd-11</i> gene during late gastrulation", <i>EMBO J.</i> 12 :3539-3550 (1993).
EG	Gurdon, J. B., "The generation of diversity and pattern in animal development", <i>Cell</i> 68 :185-199 (1992).
EH	Halpern, M. E., "Induction of muscle pioneers and floor plate is distinguished by the zebrafish <i>no tail</i> mutation", <i>Cell</i> 75 :99-111 (1993).
EI	Gustin, K. et al., "Characterization of the role of individual protein binding motifs within the hepatitis B virus enhancer 1 on X promoter activity using linker scanning mutagenesis", <i>Virology</i> 193 :653-660 (1993).
EJ	Hall, T. et al., "A potential catalytic site revealed by the 1.7-Å crystal structure of the amino-terminal signaling domain of sonic hedgehog", <i>Nature</i> 378 (6553):212-216 (Nov 1995).
EK	Hamburger, V. and H. L. Hamilton, "A series of normal stages in the development of the chick embryo", <i>J. Morph.</i> 88 :49-92 (1951).

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	EN	Hardy, A. et al., "Gene expression, polarising activity and skeletal patterning in reaggregated hind limb mesenchyme", <i>Devel.</i> 121 (12):4329-4337 (Dec. 1995).
	EO	Harmon, C. S. et al., "Evidence that activation of protein kinase A inhibits human hair follicle growth and hair fibre production in organ culture and DNA synthesis in human and mouse hair follicle organ culture", <i>British J. Dermatol.</i> 136 :853-858 (1997).
	EP	Hatta, K. et al., "The cyclops mutation blocks specification of the floor plate of the zebrafish central nervous system", <i>Nature</i> 350 :339-341 (1991).
	ER	Heberlein, U. et al., "The TGB β homolog <i>dpp</i> and the segment polarity gene <i>hedgehog</i> are required for propagation of a morphogenetic wave in the <i>Drosophila</i> retina", <i>Cell</i> 75 :913-926 (1993).
	ES	Heemskerk, J. and S. DiNardo, " <i>Drosophila hedgehog</i> acts as a morphogen in cellular patterning", <i>Cell</i> 76 :449-460 (1994).
	ET	Hidalgo, A. and P. Ingham, "Cell patterning in the <i>Drosophila</i> segment: spatial regulation of the segment polarity gene <i>patched</i> ", <i>Devel.</i> 110 :291-301 (1990).
	EU	Hooper, J. and M. Scott, "The <i>Drosophila patched</i> gene encodes a putative membrane protein required for segmental patterning", <i>Cell</i> 59 :751-765 (1989).
	EV	Hynes, R. O., "Integrins: A family of cell surface receptors", <i>Cell</i> 48 :549-554 (1987).
	EW	Hynes, R. O., "Induction of midbrain dopaminergic neurons by Sonic hedgehog", <i>Neuron</i> 15 (1):35-44 (July 1995).
	EX	Ingham, P. W., "Signaling by hedgehog family proteins in <i>Drosophila</i> and vertebrate development", <i>Curr. Opin. Genet. Dev.</i> 5 (4):478-484 (Aug 1995).
	EY	Ingham, P. W., "Hedgehog points the way", <i>Current Biology</i> 4 (4):347-350 (1994).
	EZ	Ingham, P. W., "Localized <i>Hedgehog</i> activity controls spatial limits of wingless transcription in the <i>Drosophila</i> embryo", <i>Nature</i> 366 :560-562 (1993).
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	FF	Jessel, T. M. and D. A. Melton, "Diffusible factors in vertebrate embryonic induction", <i>Cell</i> 68 :257-270 (1992).
	FG	Johnson, R. L. and C. Tabin, "The long and short of hedgehog signaling", <i>Cell</i> 81 :313-315 (5 May 1995).
	FH	Johnson, R. L. et al., "Patched overexpression alters wing disc size and pattern: transcriptional and post-transcriptional effects on hedgehog targets", <i>Devel.</i> 121 (12):4237-4245 (Dec. 1995).
	FI	Johnson, R. L. et al., "Ectopic expression of sonic hedgehog alters dorsal-ventral patterning of somites", <i>Cell</i> 79 (7):1165-1173 (Dec. 1994).
	FJ	Johnson, R. L. et al., "Mechanism of limb patterning", <i>Curr. Opin. Genet. Dev.</i> 4 (4):535-542 (Aug. 1994).
	FJ	Johnson, R. L. et al., "Sonic hedgehog: a key mediator of anterior-posterior patterning of the limb and dorso-ventral patterning of axial embryonic structures" <i>Biochem. Soc. Trans.</i> 22 (3):569-574 (Aug. 1994).



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	FM	Koonin, E., "A protein splice-junction motif in hedgehog family proteins", <i>Trends Biochem. Sci.</i> <u>20</u> (4):141-142 (April 1995).
	FN	Kornblitt, A. R. et al., "Primary structure of human fibronectin: differential splicing may generate at least 10 polypeptides from a single gene", <i>EMBO J.</i> <u>4</u> :1755-1759 (1985).
	FO	Kornfeld, R. and S. Kornfeld, "Assembly of asparagine-linked oligosaccharides", <i>Ann. Rev. Biochem.</i> <u>54</u> :631-664(1985).
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	FW	Levin, M. et al., "A molecular pathway determining left-right asymmetry in chick embryogenesis", <i>Cell</i> <u>82</u> (5):803-814 (Sept. 8, 1995).
	FX	Li, W. et al., "Function of protein kinase A in hedgehog signal transduction and drosophila imaginal disc development", <i>Cell</i> <u>80</u> (4):553-562(Feb. 1995).
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	FZ	Lumsden, A. and A. Graham, "Neural patterning: A forward role for hedgehog", <i>Curr. Biol.</i> <u>5</u> (12):1347-1350 (Dec. 1995).
	GA	Ma, C. et al., "Molecular cloning and characterization of rKik10, a cDNA encoding T-kininogenase from rat submandibular gland and kidney", <i>Biochem.</i> <u>31</u> (44):10922-10928 (1992).
	GB	Ma, C. et al., "The segment polarity gene <i>hedgehog</i> is required for the progression of the morphogenetic furrow in the developing Drosophila eye", <i>Cell</i> <u>75</u> :927-938 (1993).
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	GE	Maccabe, J. A. and B. W. Parker, "The target tissue of limb-bud polarizing activity in the induction of supernumerary structures", <i>J. Embryol. Exp. Morph.</i> <u>53</u> :67-73 (1979).
	GF	Maiese, K. et al., "Protein kinases modulate the sensitivity of hippocampal neurons to nitric oxide toxicity and anoxia", <i>J. Neurosci. Res.</i> <u>36</u> :77-87 (1993).
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	GI	Mavilio, F. et al. "Activation of four homeobox gene clusters in human embryonal carcinoma cells induced to differentiate by retinoic acid", <i>Differentiation</i> <u>37</u> :73-79 (1988).

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	GL	Mohler, J. and K. Vani, "Molecular organization and embryonic expression of the <i>hedgehog</i> gene involved in cell-cell communication in segmental patterning of <i>Drosophila</i> ", <i>Devel.</i> 115 :957-971 (1992).
	GM	Morgan, B. A. et al., "Targeted misexpression of <i>Hox-4.6</i> in the avian limb bud causes apparent homeotic transformations", <i>Nature</i> 358 :236-239 (1992).
	GN	Nakano, Y. et al., "A protein with several possible membrane-spanning domains encoded by the <i>Drosophila</i> segment polarity gene <i>patched</i> ", <i>Nature</i> 341 :508-513 (1989).
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	GP	Niswander, L. and G. R. Martin, "FGF-4 and BMP-2 have opposite effects on limb growth", <i>Nature</i> 361 :68-71(1993).
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	GR	Nohno, T. et al., "Involvement of the <i>Chox-4</i> Chicken Homeobox Genes in Determination of Anteroposterior Axial Polarity during Limb Development", <i>Cell</i> , Vol. 64 : 1197- 1205 (March 22, 1991).
	GS	Nohno, T. et al., "Involvement of the Sonic hedgehog gene in chick feather formation", <i>Biochem. Biophys. Res. Comm.</i> 206 (1): 33-39 (Jan. 1995).
	GT	O'Farrell, P. H., "Unanimity waits in the wings", <i>Nature</i> 368 :188-189 (1994).
	GU	Parisi, M. J. et al., "The role of the hedgehog/patched signaling pathway in epithelial stem cell proliferation: From fly to human", <i>Cell Res.</i> 8 :15-21 (1998).
	GV	Parr, B. A. et al., "Mouse Wnt genes exhibit discrete domains of expression in the early embryonic CNS and limb buds", <i>Development</i> 119 :247-261 (1993).
	GW	Patel, N. H. et al., "The role of segment polarity genes during <i>Drosophila</i> neurogenesis", <i>Genes & Devel.</i> 3 :890-904 (1989).
	GX	Peifer, M., "The two faces of hedgehog", <i>Science</i> 266 (5190):1492-1493 (Dec. 1994).
	GY	Perrimon, N. et al., "Generating lineage-specific markers to study <i>Drosophila</i> development", <i>Develop. Genet.</i> , 12 :238-252 (1991).
	GZ	Perrimon, N., "Hedgehog and beyond", <i>Cell</i> 80 :517-520 (24 Feb. 1995).
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	HB	Phillis, J. W. and M. H. O'Regan, "Mechanisms of glutamate and aspartate release in the ischemic rat cerebral cortex", <i>Brain Res.</i> 730 :150-164 (1996).
	HC	Placzek, M. et al., "Induction of floor plate differentiation by contact-dependent, homeogenetic signals", <i>Development</i> 117 : 205-218 (1993).
	HD	Placzek, M. et al., "Orientation of Commissural Axons <i>in vitro</i> in response to a floor plate-derived chemoattractant", <i>Develop.</i> 110 :19-30 (1990).
	HE	Pollock, R. A. et al., "Altering the boundaries of <i>Hox3.1</i> expression: Evidence for antipodal gene regulation", <i>Cell</i> 71 :911-923 (1992).
	HF	Porter, J. et al., "The product of hedgehog autoproteolytic cleavage active in local and long-range signalling", <i>Nature</i> 374 (6520):363-366 (23 March 1995).
	HG	Reeck, et al., "'Homology' in proteins and nucleic acids: A terminology muddle and a way out of it", <i>Cell</i> 50 :667 (28 Aug. 1987).
✓	HH	Rennie, J., "Super Sonic", <i>Sci. Amer.</i> p.20, (April 1994).

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HI	Riddle, R. D. et al., "Sonic hedgehog mediates the polarizing activity of the ZPA", <i>Cell</i> 75 :1401-1416, (31 Dec. 1993).
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EXAMINER	DATE CONSIDERED
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